

आज अगर आप 6 घंटे पढ़ने के लिए तैयार नहीं हो सकते
तो कल आपको 12 घंटे काम करने के लिए तैयार रहना होगा!

CSIR NET – Life Science

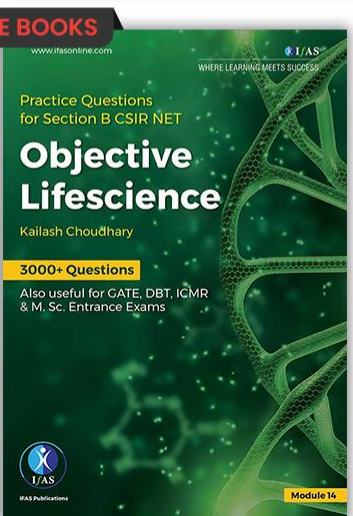
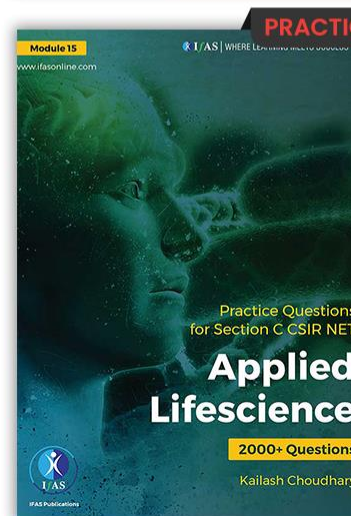
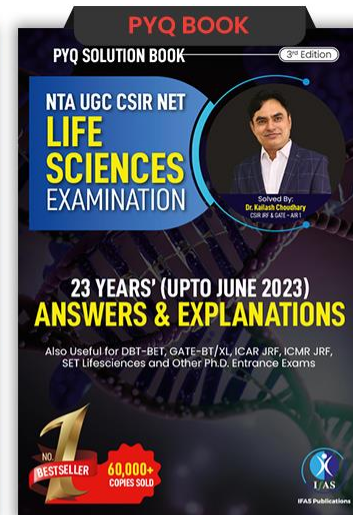
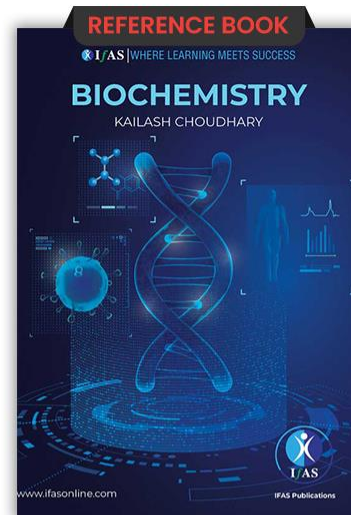
Unit 1: Biochemistry

07

Lipids and Nucleotides



Order Online and Get
Free Delivery Across India





Points to be covered in this Lecture



What are Lipids?



Fatty Acids



Simple Lipids



Glycerol Based Lipids



Sphingosine Based Lipids



Cholesterol



Nitrogenous Bases



Nucleoside and Nucleotides



Lipids are esters of fatty acids and polyhydroxy alcohol.

Simple lipids are the compounds of C, H, O but the ratio of H and O is more than 2:1

$$\begin{matrix} C : H >> O \\ \underline{1} : \underline{2} \end{matrix}$$

Their general formula is $\text{C}_n\text{H}_{2n}\text{O}_2$.

- Non-polar
- Macromolecule
- Not true polymer



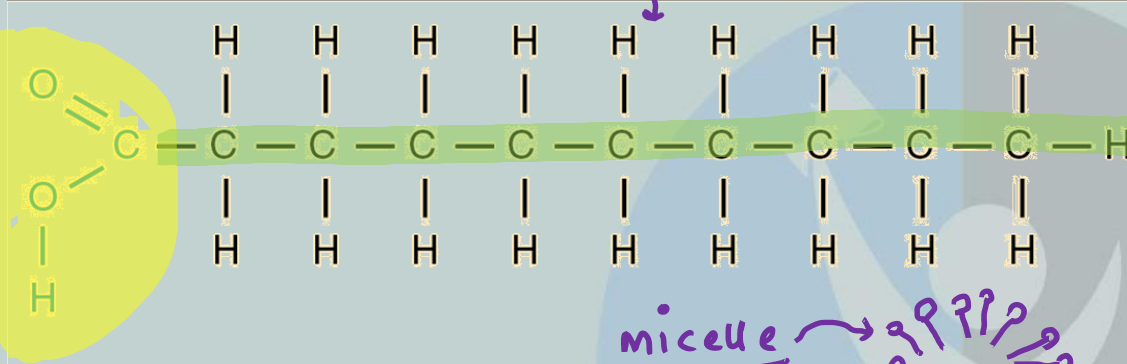
Preferred Energy Storage molecule:

- Lipids represent highly reduced forms of carbon and,
- Lipids upon oxidation yield large amounts of energy.
- Lipids are choice for metabolic energy storage.
- Preferred for marathon runner
- Lipid are composed of
 - Fatty Acids
 - Polyhydroxy Alcohol

• Carbohydrate $4 \frac{\text{K-cal}}{\text{g}}$
• Protein 4.2
• Lipids $9 \frac{\text{K-cal}}{\text{g}}$
* occupy lesser space

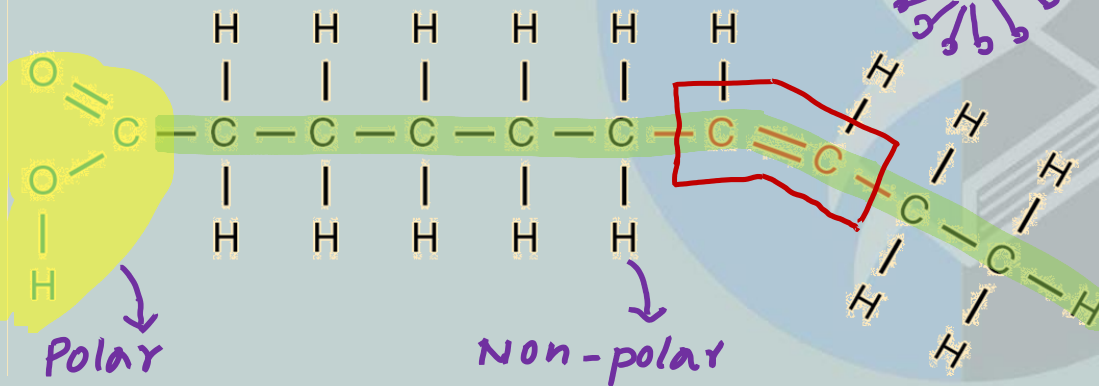


I. Fatty Acids: Aliphatic chain with Acidic Group



Saturated Fatty Acid

- Only single bonds
- Completely Reduced
- Yield more energy



Unsaturated Fatty Acid

- 1 or more double bond
- Partially oxidized.

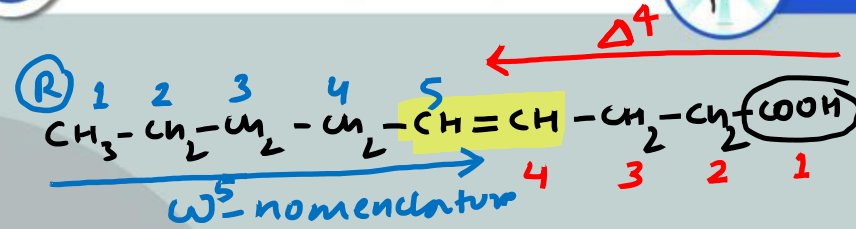


Saturated fatty acids

Formula		Common Name	Melting Point
$\text{CH}_3(\text{CH}_2)_{10}\text{COOH}$	12C	Lauric acid	45°C
$\text{CH}_3(\text{CH}_2)_{12}\text{COOH}$	14C	Myristic acid	55°C
$\text{CH}_3(\text{CH}_2)_{14}\text{COOH}$	16C	<u>Palmitic acid</u>	63°C
$\text{CH}_3(\text{CH}_2)_{16}\text{COOH}$	18C	Stearic acid	69°C
$\text{CH}_3(\text{CH}_2)_{18}\text{COOH}$	20C	Arachidic acid	76°C

✓ Unsaturated fatty acids

<u>16:1</u> <u>Cis-Δ^9</u>	<u>ω^7</u>	<u>Palmitoleic acid</u>	0°C
<u>18:1</u> <u>Cis-Δ^9</u>	<u>ω^9</u>	<u>Oleic acid</u>	13°C
<u>18:1</u> <u>Trans-Δ^9</u>	<u>ω^9</u>	Elaidic acid	45°C
<u>18:2</u> <u>Cis-$\Delta^{9,12}$</u>	<u>ω^6</u>	<u>Linoleic acid</u>	-5°C
<u>18:3</u> <u>Cis-$\Delta^{6,9,12}$</u>	<u>ω^6</u>	γ -Linolenic acid	-11°C
<u>18:3</u> <u>Cis-$\Delta^{9,12,15}$</u>	<u>ω^3</u>	α -Linolenic acid	-17°C
<u>20:4</u> <u>Cis-$\Delta^{5,8,11,14}$</u>	<u>ω^6</u>	Arachidonic acid	-49°C
<u>20:5</u> <u>Cis-$\Delta^{5,8,11,14,17}$</u>	<u>ω^3</u>	Timnodonic acid	-69°C
<u>22:6</u> <u>Cis-$\Delta^{4,7,10,13,16,19}$</u>	<u>ω^3</u>	Cervonic acid	-92°C



Saturated FA

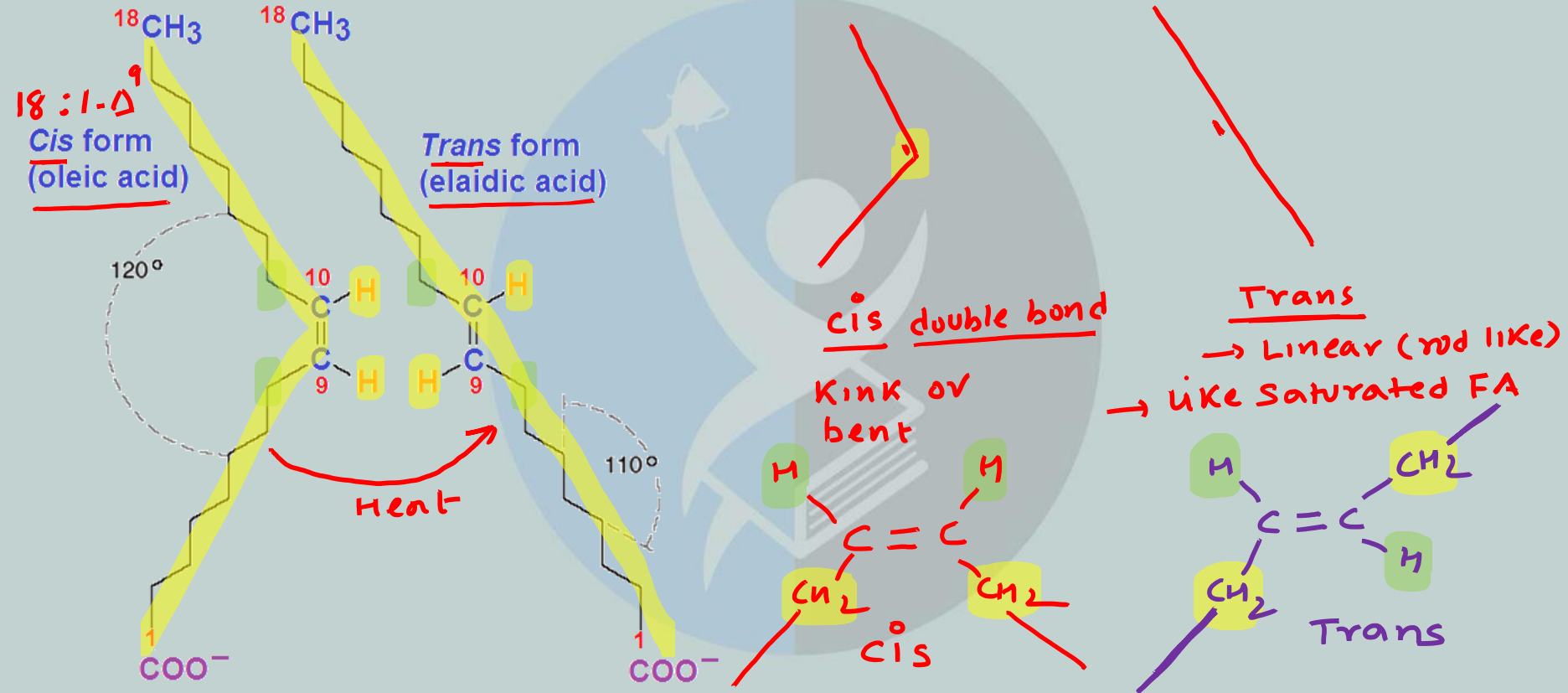
Mono unsaturated FA (MUFA)

Non-essential FA

Poly-unsaturated FA
(PUFA)

Essential FA (ω^3 & ω^6)

Cis-double bonds are more preferred in naturally occurring unsaturated fatty acids



Saturated fatty acids

Formula		Common Name	Melting Point
$\text{CH}_3(\text{CH}_2)_{10}\text{COOH}$	12C	Lauric acid	45 °C
$\text{CH}_3(\text{CH}_2)_{12}\text{COOH}$	14C	Myristic acid	55 °C
$\text{CH}_3(\text{CH}_2)_{14}\text{COOH}$	16C	Palmitic acid	63 °C
$\text{CH}_3(\text{CH}_2)_{16}\text{COOH}$	18C	Stearic acid	69 °C
$\text{CH}_3(\text{CH}_2)_{18}\text{COOH}$	20C	Arachidic acid	76 °C

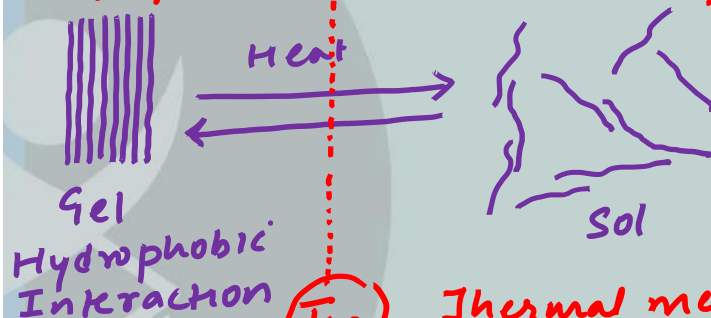
Unsaturated fatty acids

16:1 Cis- Δ^9	ω^7	Palmitoleic acid	0 °C
18:1 Cis- Δ^9	ω^9	Oleic acid	13 °C
18:1 Trans- Δ^9	ω^9	Elaidic acid	45 °C
18:2 Cis- $\Delta^{9,12}$	ω^9	Linoleic acid	- 5 °C
18:3 Cis- $\Delta^{6,9,12}$	ω^6	γ - Linolenic acid	- 11 °C
18:3 Cis- $\Delta^{9,12,15}$	ω^3	α - Linolenic acid	- 17 °C
20:4 Cis- $\Delta^{5,8,11,14}$	ω^6	Arachidonic acid	- 49 °C
20:5 Cis- $\Delta^{5,8,11,14,17}$	ω^3	Timnodonic acid	- 69 °C
22:6 Cis- $\Delta^{4,7,10,13,16,19}$	ω^3	Cervonic acid	- 92 °C

The melting point (T_m) of fatty acids,

- ① increases with chain length and
- ② decreases with the degree of (cis)-unsaturation.

Increase
50%.



50%.

Thermal melting point

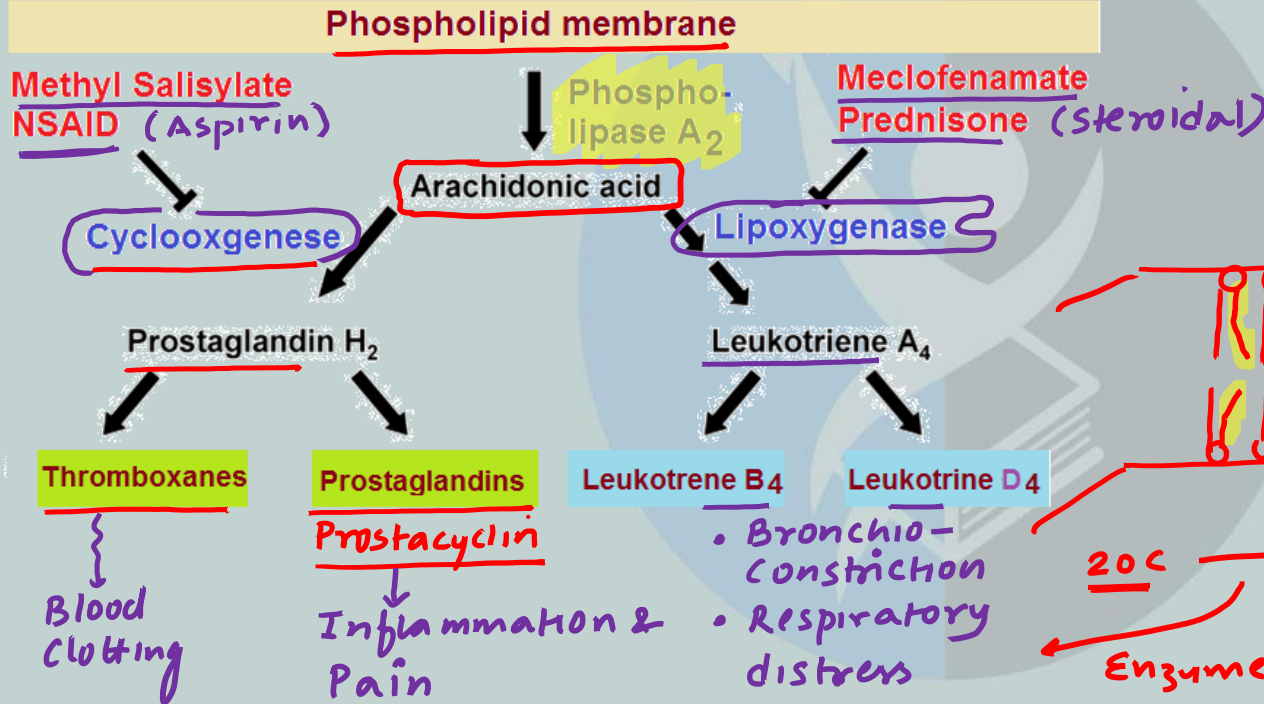
$$\Delta G = 0 \quad T_m = \frac{\Delta H}{\Delta S}$$

Decrease



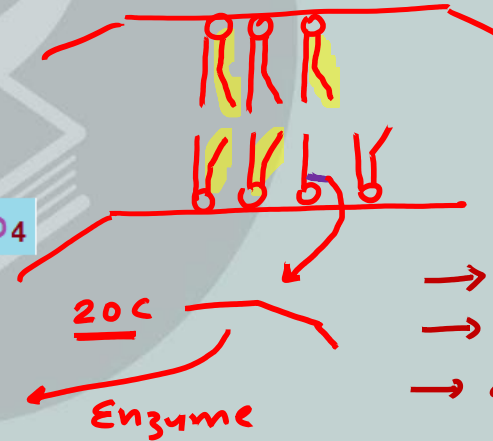
Eicosanoids: 20 carbon derivative.

Arachidonic acid, 20 carbon units derivative



✓ PUFA are essential

- cannot be synthesized by animals but are **important** for various **physiological function**.
- obtained from diet



- Thromboxane
- Prostaglandin
- Prostacyclin
- Leukotriene

Eicosanoids

- Local hormone
- Half life - Low
- GPCR family receptors

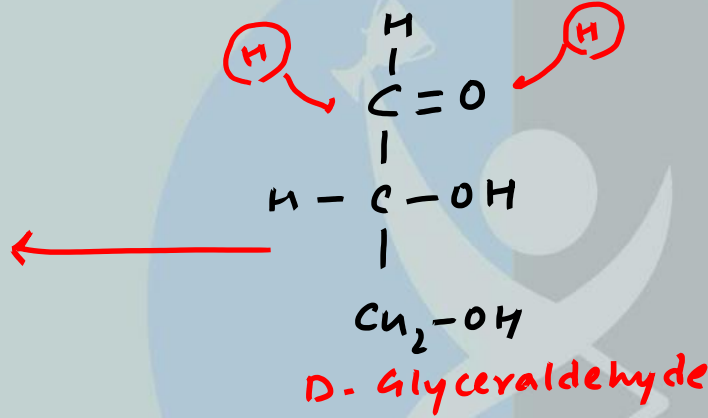
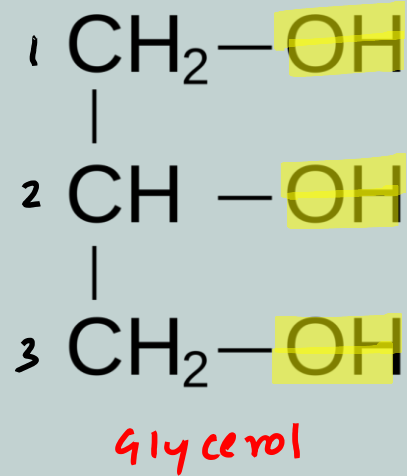
20C

Enzyme

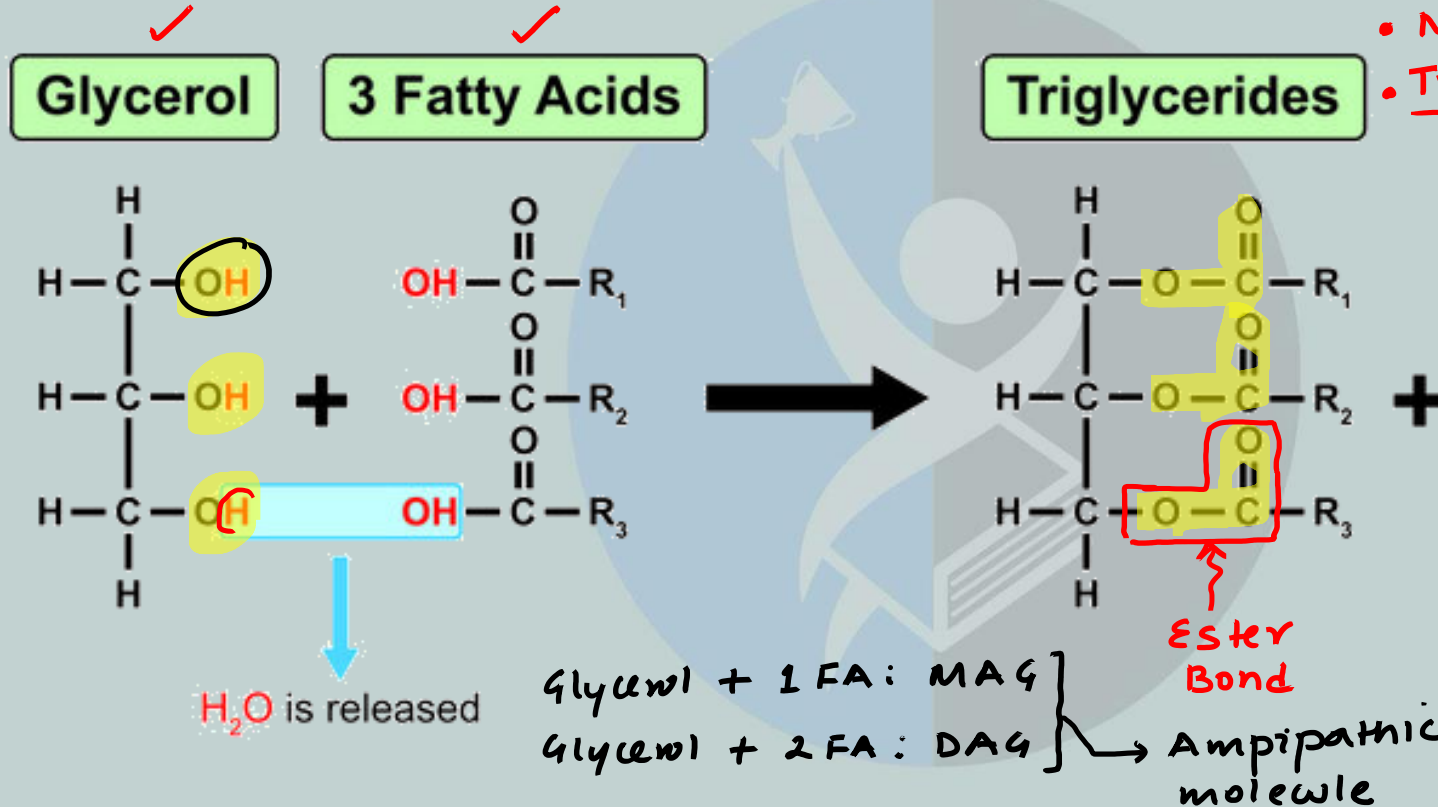


Polyhydroxy Alcohol- Glycerol : Polar

Derived from reduction of Glyceraldehyde



Fats and oils (Tri-Acyl Glycerol): Ester of Fatty Acid and Glycerol



- Do not form membrane
- Non-polar.

- Tri-Acyl Glycerol (TAG)
- Energy storage form

Fat
 ↓
 Animals

oil
 ↓
 plant

Ester Bond



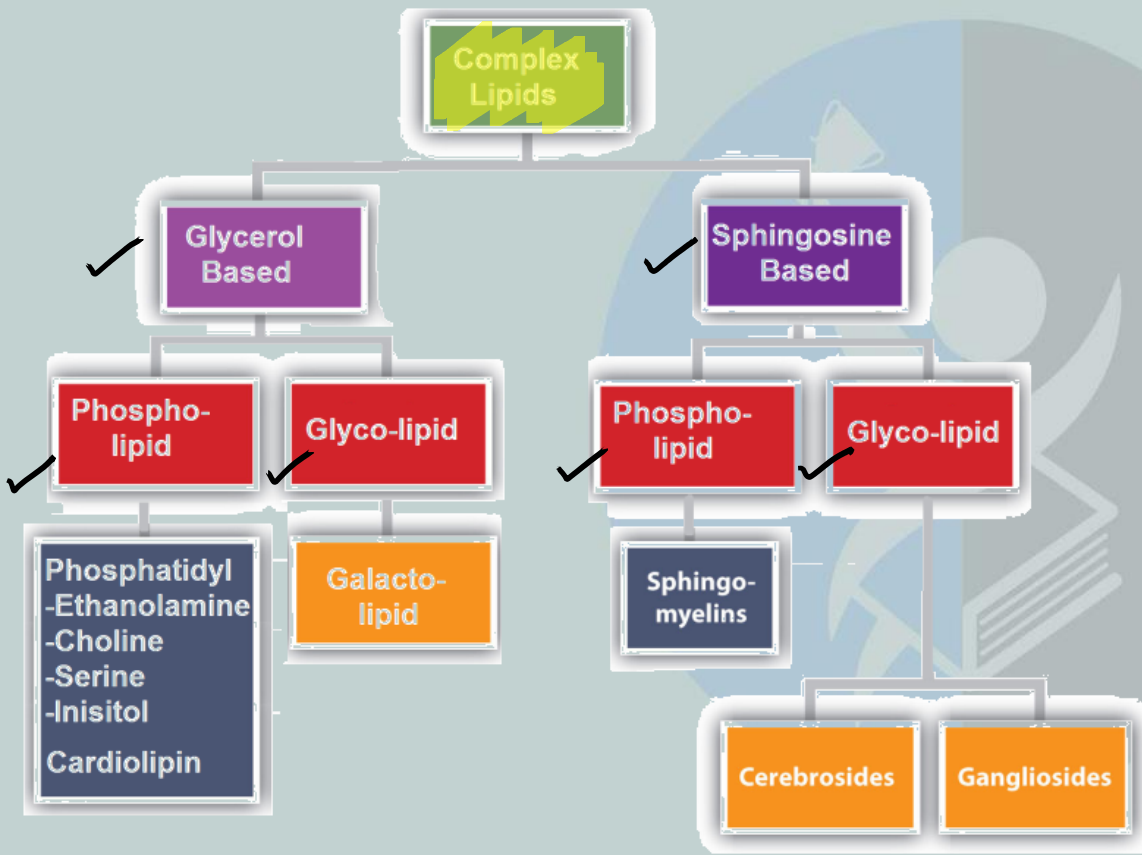
Membrane lipids are derived Lipids



+
Extra group

Phosphate

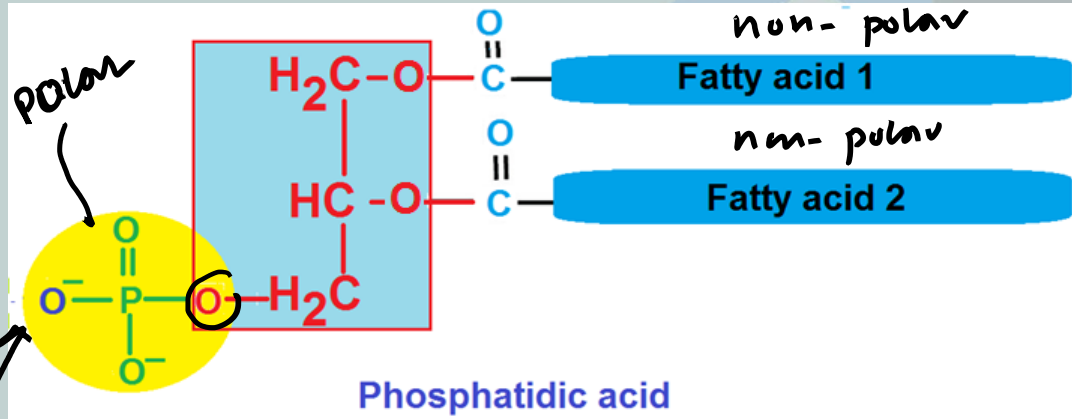
Sugar

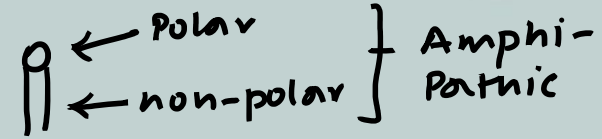




A. Glycerophospholipids

Derivatives of phosphatidic acid.



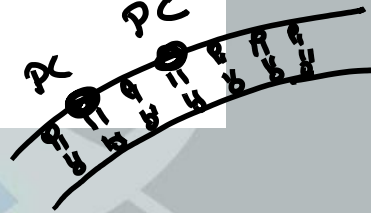
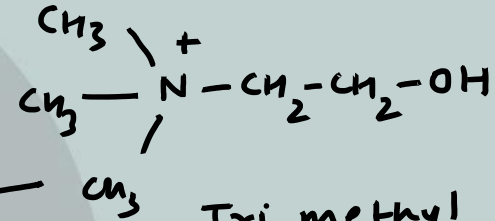
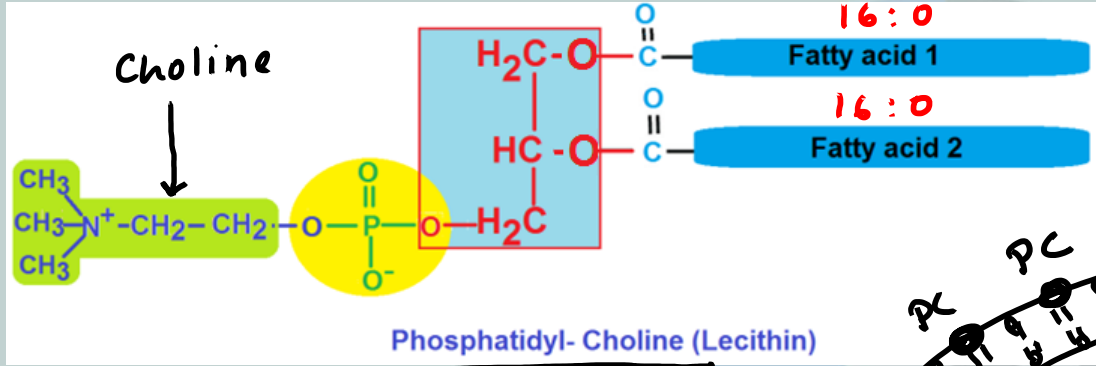

 ← Polar
 ← non-polar } Amphipathic

Synthesis: SER in eukaryotes or plasma membrane of bacteria

1 glycerol + 2 FA + 1 phosphate



Phosphatidyl-choline (lecithin): 1 choline + 1 phosphate + 1 Glycerol + 2 FA



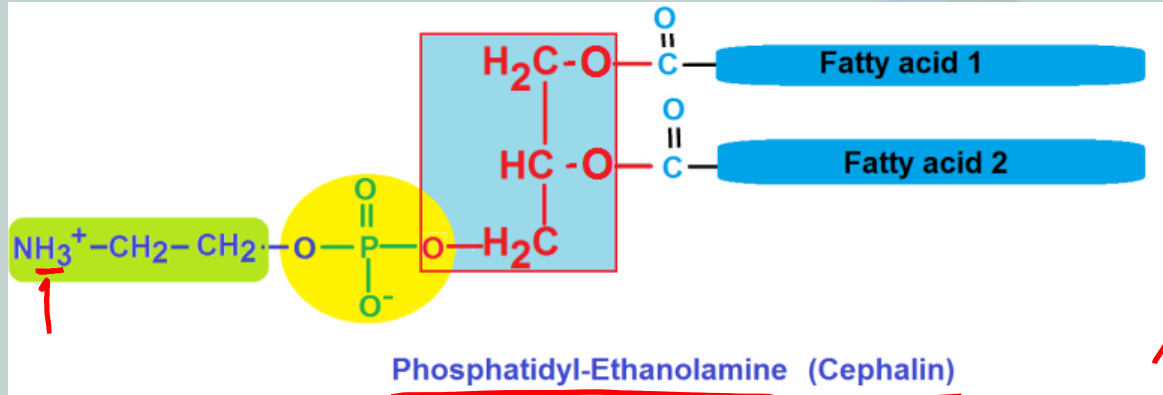
Dipalmitoyl Phosphatidyl Choline (DPPC)

Dipalmitoyl lecithin: surfactant

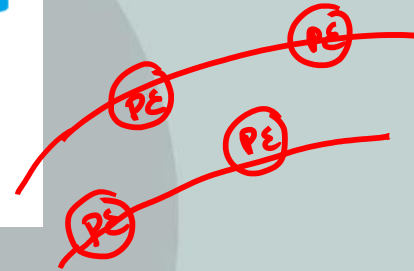
- ✓ Produced by Type II alveolar epithelial cells (pneumocytes) ← lung alveoli
- ✓ Absence of surfactant leads to Respiratory distress syndrome.



Phosphatidylethanolamine (cephalin): 1 ethanolamine + 1 phosphate
+ 1 glycerol
+ 2 Fatty acid

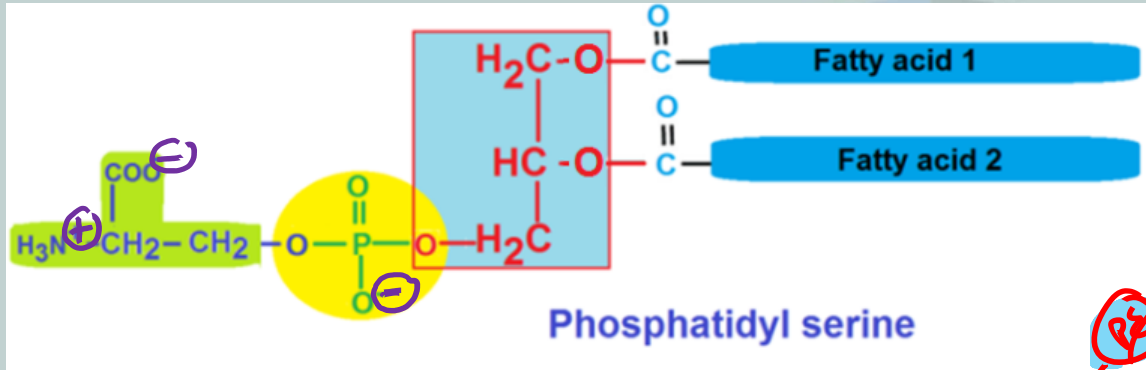


• Abundant in bacterial membrane





Phosphatidyl serine: 1 serine + 1 phosphate + 1 glycerol + 2 FA



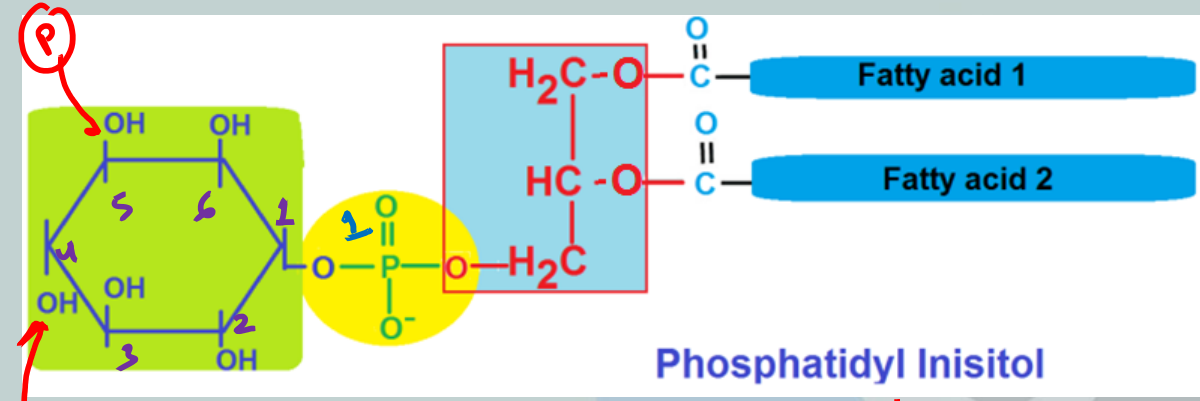
- present on cytosolic face of membrane
- PS if on outside — eat me signal
— Apoptotic cell.

→ PS can be recognized by Annexin-V protein

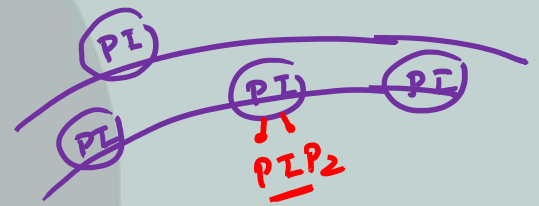




Phosphatidylinositol: 1 Inositol + 1 Phosphate + 1 Glycerol + 2 FA



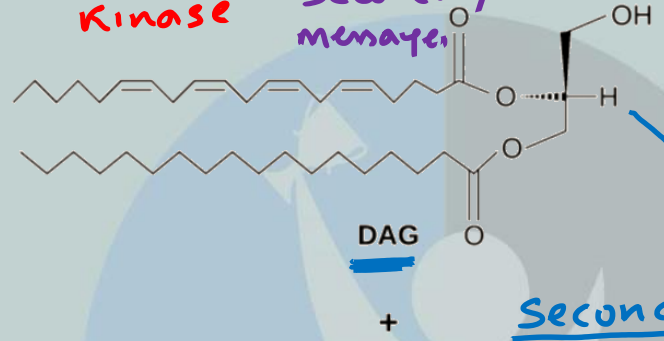
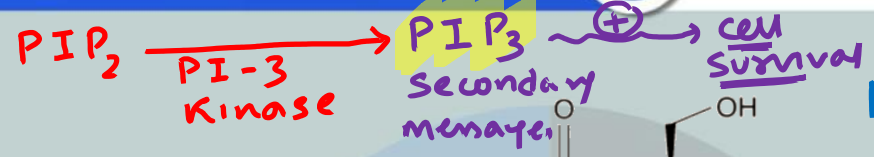
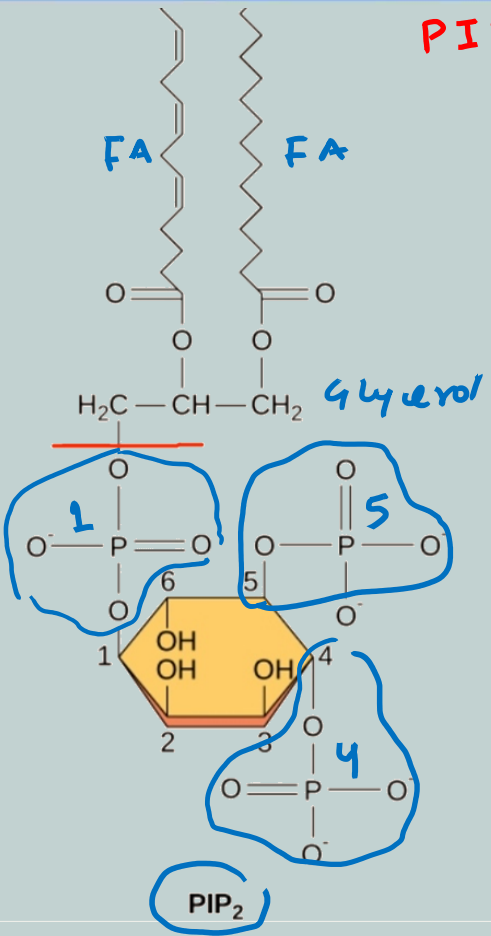
• Present more on cytosolic face



ATP → Add Phosphate Group on 4th & 5th Position of Inositol
 ↓
 ADP

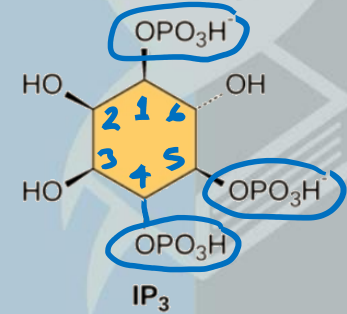
← PI-4 Kinase
 ← PI-5 Kinase

Phosphatidylinositol 4,5 Bis Phosphate (PIP₂)



activate Protein Kinase C along with Ca²⁺ ions.

Phospholipase C



opens ligand (IP₃) gated Ca²⁺ channel of ER.

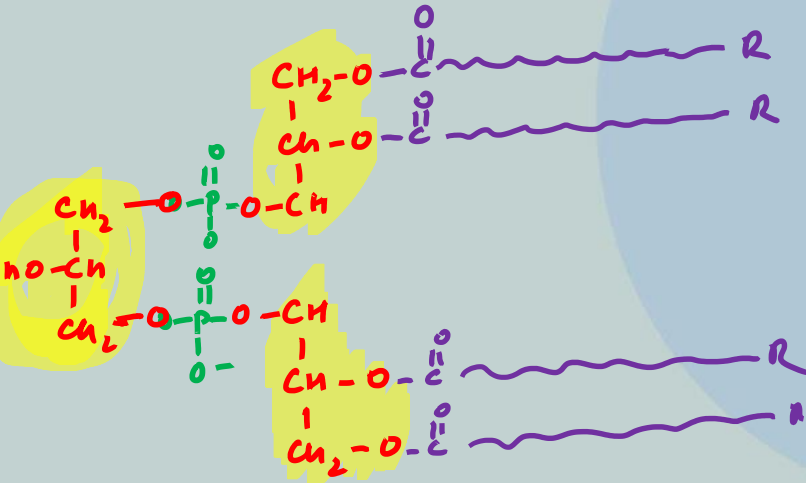
Inositol 1,4,5 Tris Phosphate (IP₃)



Cardiolipin (Diphenosphatidylglycerol):

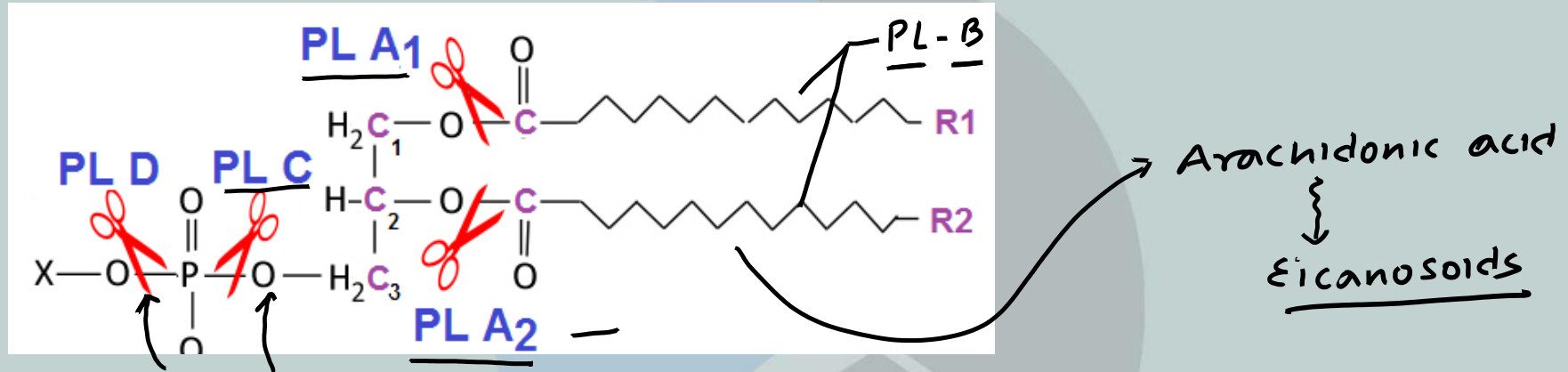
- Unique membrane lipid of inner mitochondrial membrane and bacterial plasma membrane.
- 3 Glycerol + 2 phosphate + 4 Fatty acid

• Diphenosphatidyl Glycerol



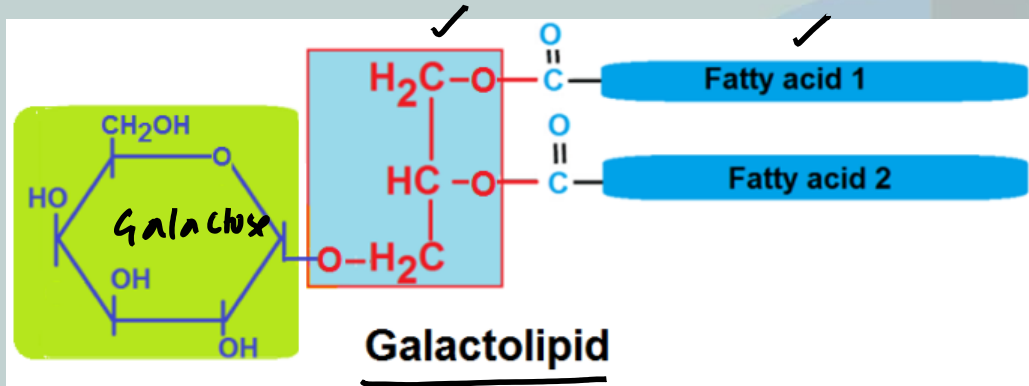


Rattlesnake and the Indian cobra (*Naja naja*) : Phospholipase A2,





B. Glycerol based glycolipids:



- 1 Galactose +
- 1 glycerol +
- 2 FA

↓

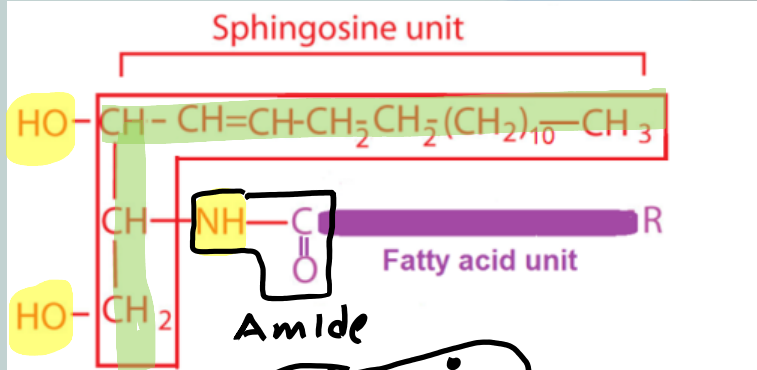
• Galactolipid

- ↳ Absent in plasma membrane
- ↳ Thylakoid membrane of chloroplast.
 - Galactolipid
 - Sulpholipids.



C. Phospho-sphingolipids:

Sphingosine : 18 carbon poly hydroxy amino alcohol.



1 Sphingosine + 1 FA

{ Amide

ceramide

(Resembles DAG)

ceramide

• Synthesis in SER

addition occurs in Golgi

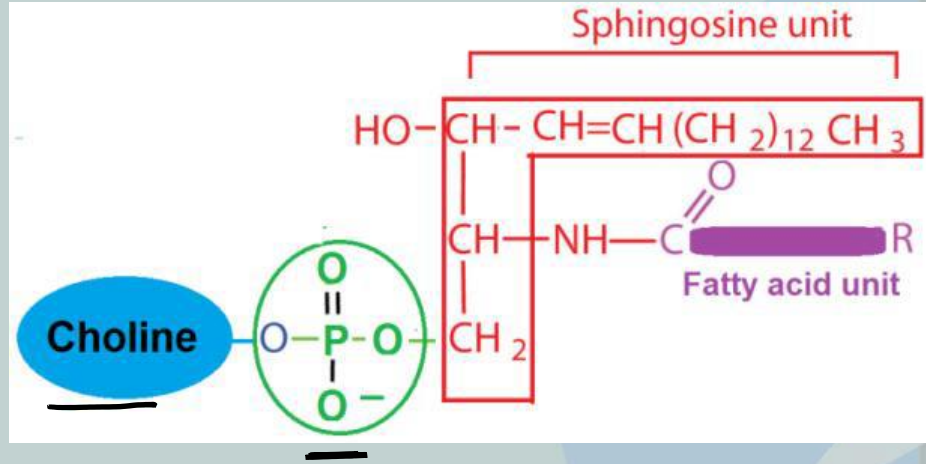
Sphingolipids

- Eukaryotes only
- only on extracellular side

Polar Group
Phospho sugar



Sphingomyelins — Phospho - Sphingo lipid



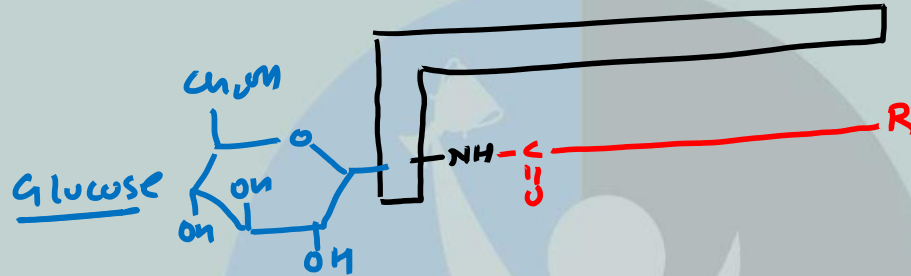
1 choline + 1 phosphate +
1 sphingosine + 1 FA

- Abundant in neural membrane & myelin sheath.
- exoplasmic face of membrane



D. Glyco-sphingolipids:

Cerebroside:



ceramide + monosaccharide : cerebroside

eg → Gluco-cerebroside
Galacto-cerebroside

Globoside: A **globoside** is a type of glycosphingolipid with more than one sugar

oligosaccharide

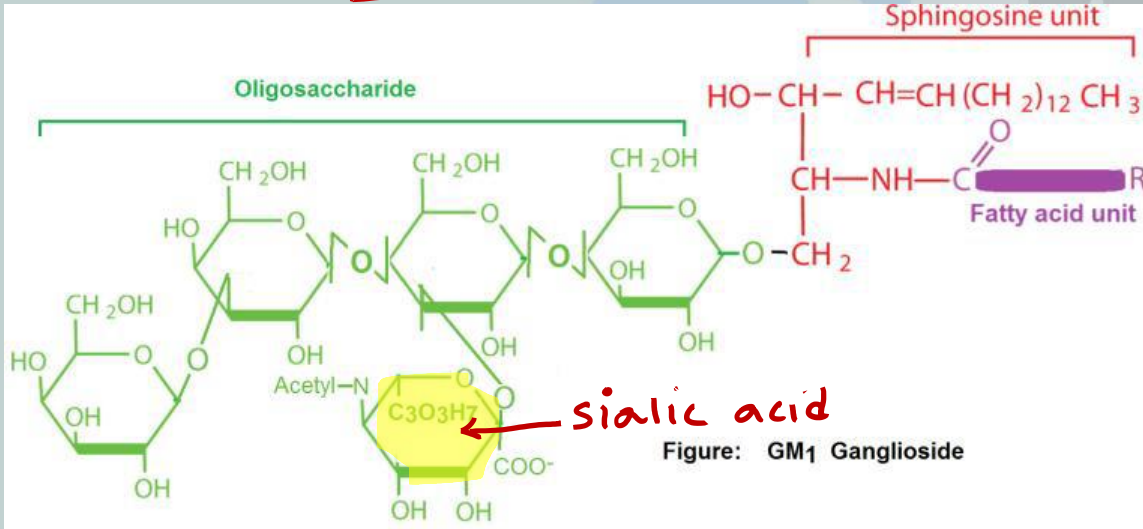
Gangliosides are complex **glycosphingolipids** derived from glucosylceramide that contain in addition one or more molecules of a **sialic acid**. (N-acetyl-D-Neuraminic acid)

GM - monosialate

GM₁ Ganglioside: Receptor for cholera toxin

GM₂ Ganglioside: accumulates in Tay-Sach disease

GM₃ Ganglioside: Receptor for influenza virus

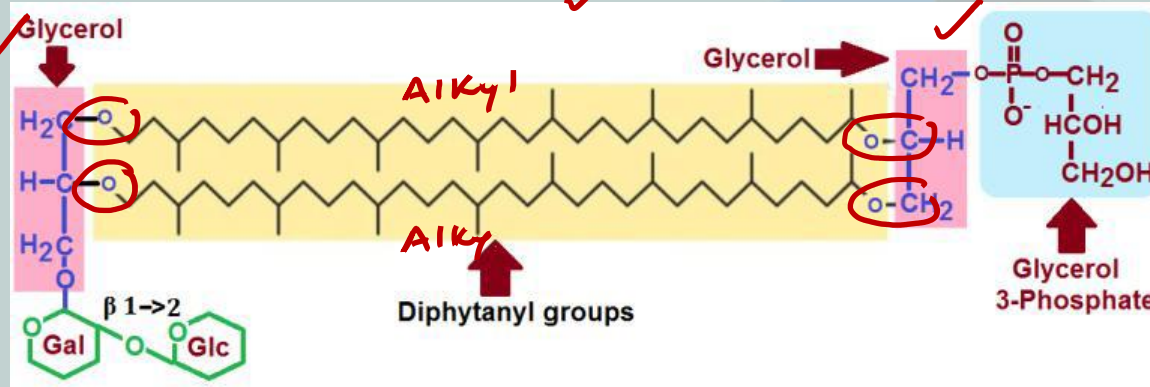


sialic acid

Figure: GM₁ Ganglioside



Archaeobacterial glycerol dialkyl glycerol tetraether:



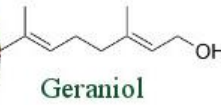
- Branched aliphatic chain
- Ether bond.
- L- Glycerol pro- chiral form.



Terpenoids

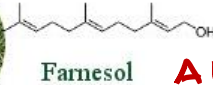
Monoterpenes

10C



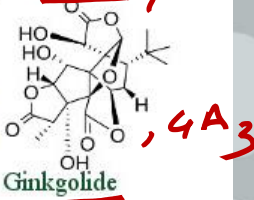
Sesquiterpenes

15 C



Diterpenes

20C



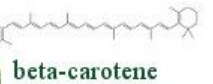
Triterpenes
30C

30c



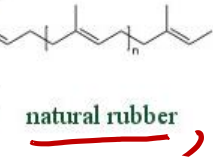
Tetraterpenes

40C



Polyterpenes

> 400

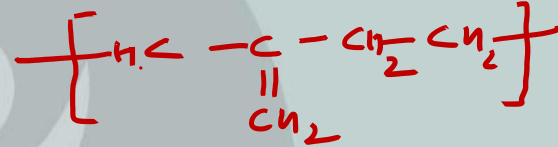


Terpene:

Terpene:

- Primary : ADA, 40%, steroid
- Secondary : Rubber, Steroids

Formed from combinations of two or more molecules isoprene (a five-carbon unit).



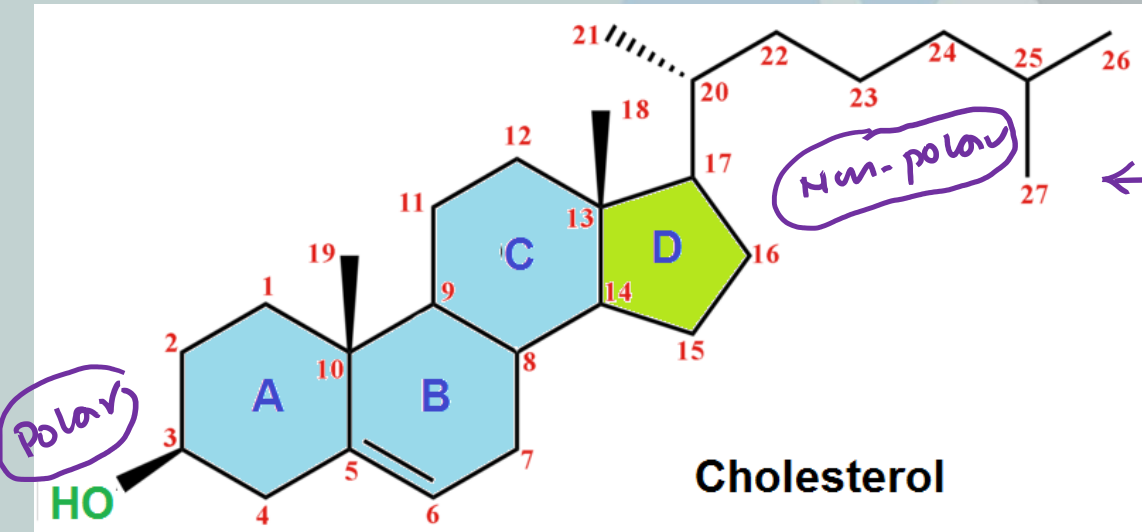
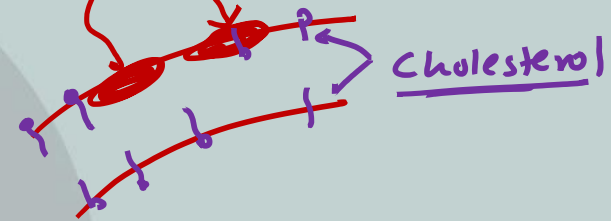
new, steroids

Quinone
Dolichol.



Cholesterol : *unique sterol of animals derived from squalene (triterpene)*

- ✓ Principal component of animal cell plasma membranes (lipid rafts). + sphingolipids
- ✓ Provides **rigidity**
- ✓ **Temperature buffer.**
- ✓ It donot influence Tm → depends on (FA)



← Amphipathic molecule



Lipid storage diseases

Autosomal Recessive
Genetic Disorder
Gene mutation
Enzyme absent

← Accumulation
in Lysosome

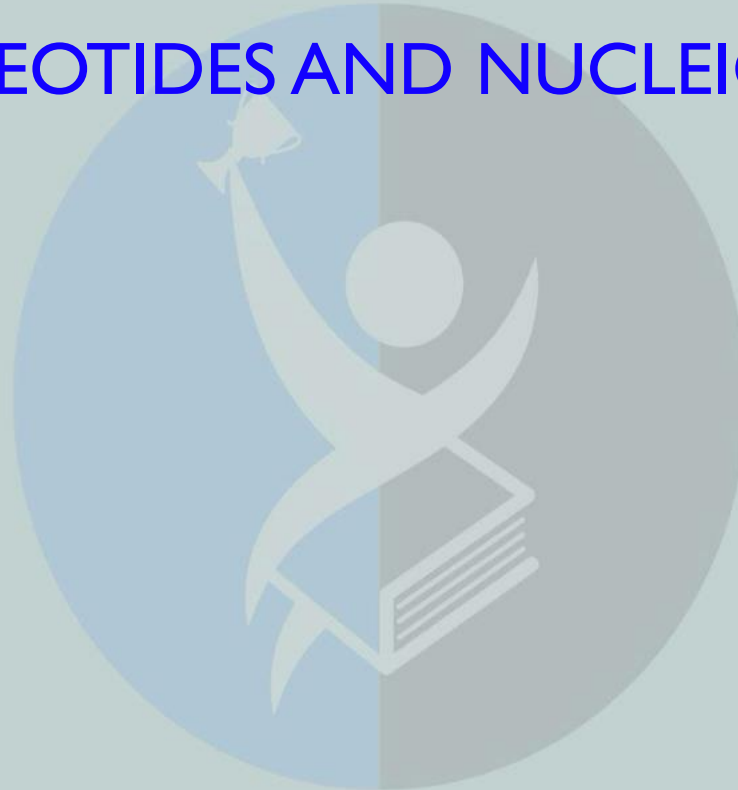
1. Nieman-pick disease: Sphingomyelinase

2. Gaucher's disease: Glucocerebrosidase

3. Tay-Sach's disease: Hexoseaminidase or
G_M₂ - Gangliosidase



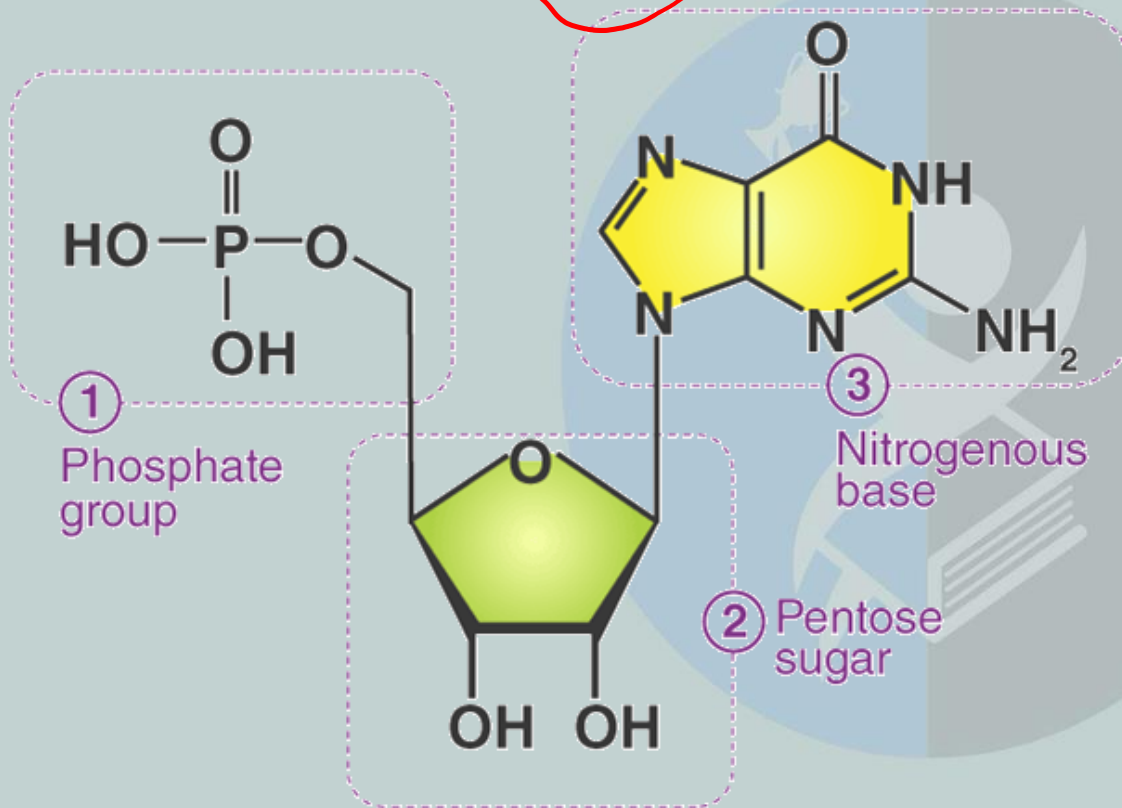
NUCLEOTIDES AND NUCLEIC ACIDS





What are nucleotides?

At least 1 Phosphate + 1 Sugar + 1 Base



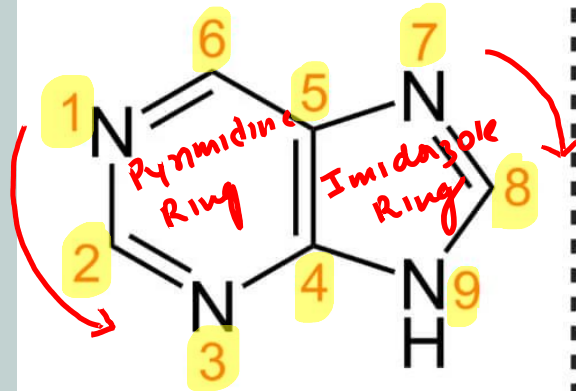
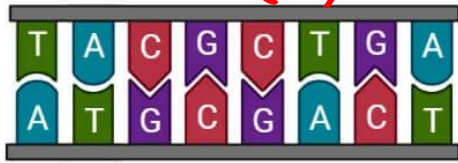


What are functions of nucleotides?

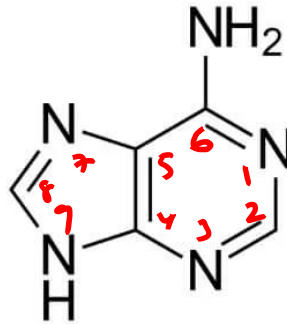
- ✓ 1. Building blocks of nucleic acids (DNA and RNA).
- ✓ 2. Involved in energy storage, muscle contraction, active transport, maintenance of ion gradients – ATP, GTP, CTP, UTP.
- ✓ 3. Activated intermediates in biosynthesis (e.g. UDP-glucose, S-adenosylmethionine).
Glycogen
↑
Ethylene
↑
- ✓ 4. Components of coenzymes (NAD⁺, NADP⁺, FAD, ~~FMN~~, and CoA)
5. Metabolic regulators:
 - ✓ • Second messengers (cAMP, cGMP)
 - ✓ • Phosphate donors in signal transduction (ATP)
 - ✓ • Regulation of some enzymes via adenylation and uridylylation

Nitrogenous Bases

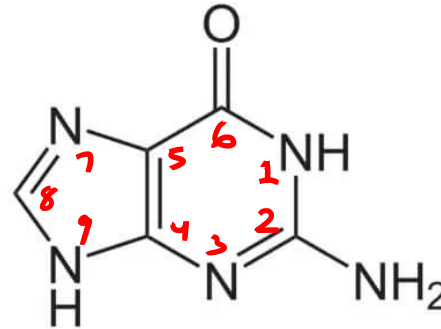
I. Purine : Double heterocyclic Ring, weak base, non-polar
(R)



Purine



Adenine

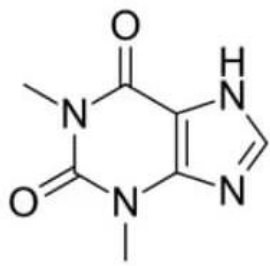
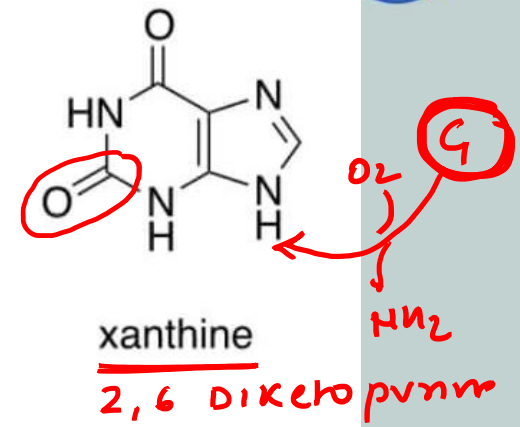
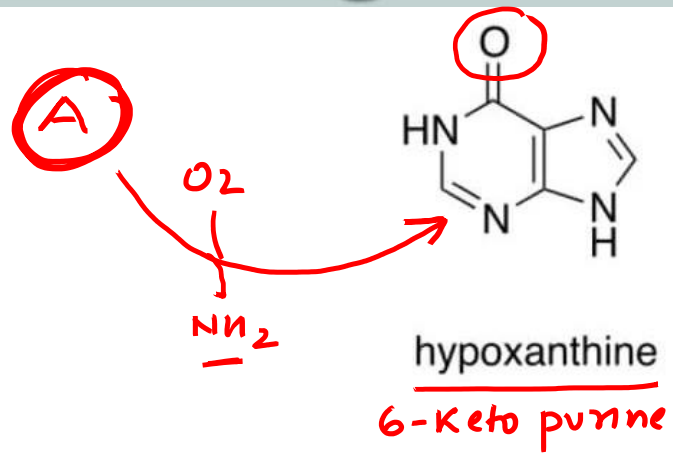


Guanine

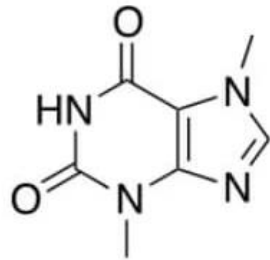
(A) 6-Amino Purine
(G): 2-Amino 6-keto purine
(R) Purine
(A) (G)



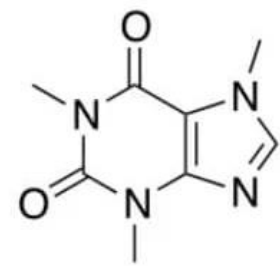
Purine Derivatives



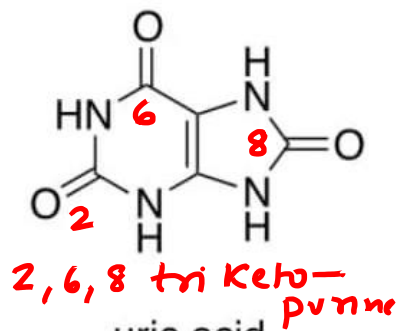
theophylline



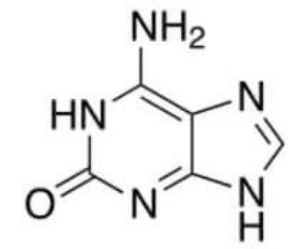
theobromine



caffeine



uric acid



isoguanine

Pyrimidine (Y)



✓ Thymine

(DNA ~~only~~)

Rare in RNA

common in



✓ Cytosine

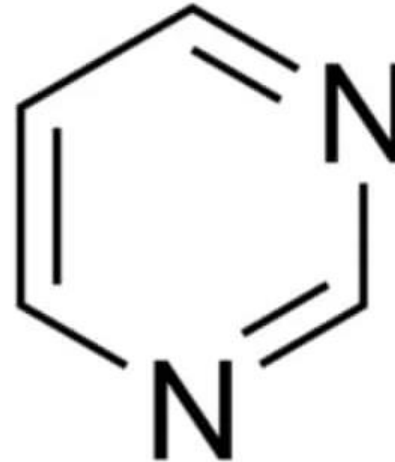


✓ Uracil

(RNA ~~only~~)

• Rare in DNA

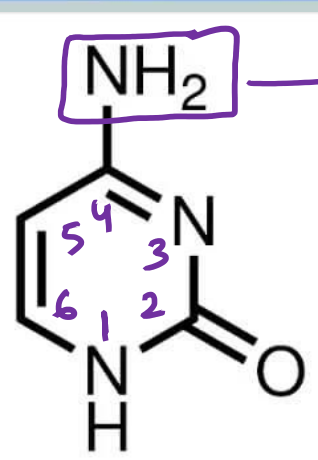
common



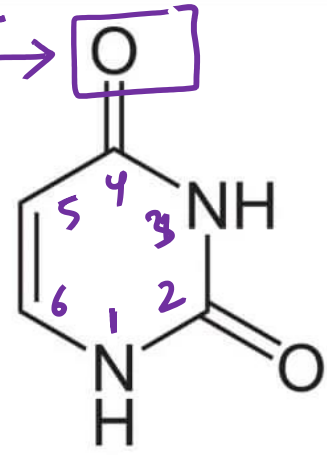
(4C & 2N)

← Diazine Ring

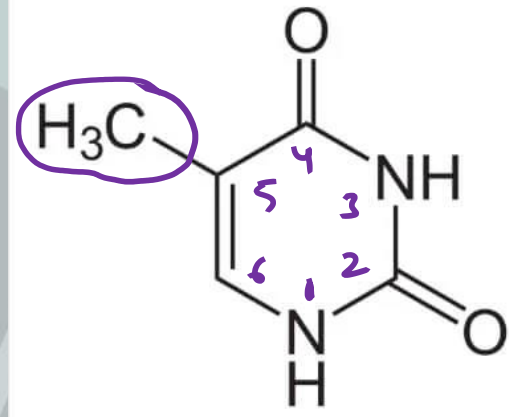
Pyrimidine
Structure



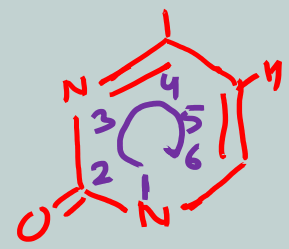
Cytosine
(Amino pyrimidine)



Uracil

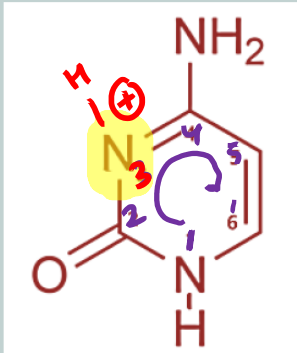
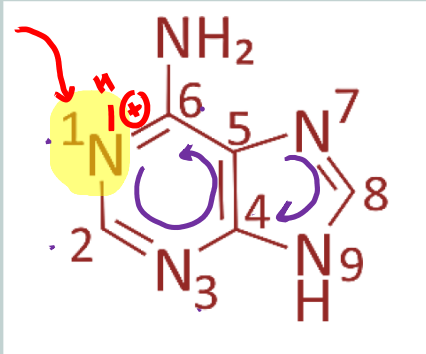


Thymine
(5-methyl uracil)





Site of Protonation



Strong acidic condition ($\text{pH} \approx 3$)

Purine : N 1

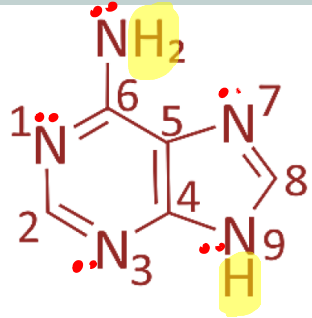
Pyrimidine : N 3



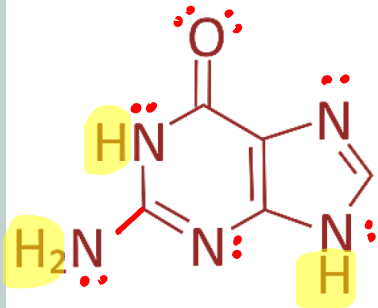
Maximum Hydrogen bond capability



- Guanine
 - ↳ various lone pairs
 - ↳ H-Donor group.

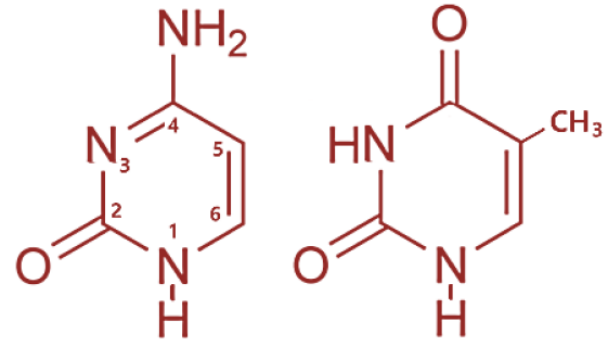


adenine



guanine

A = T or A = U
G = C

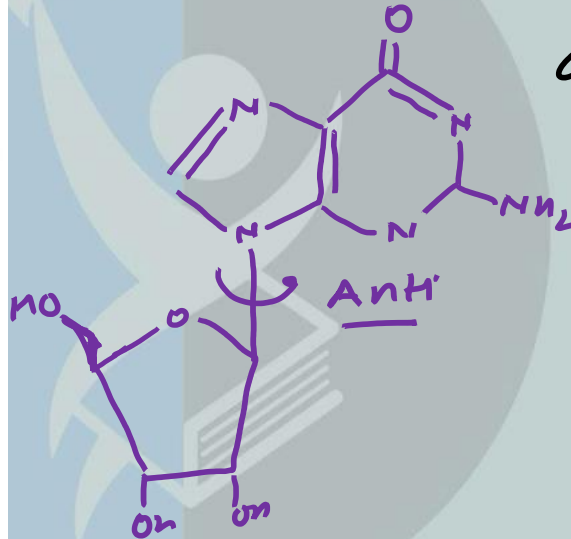
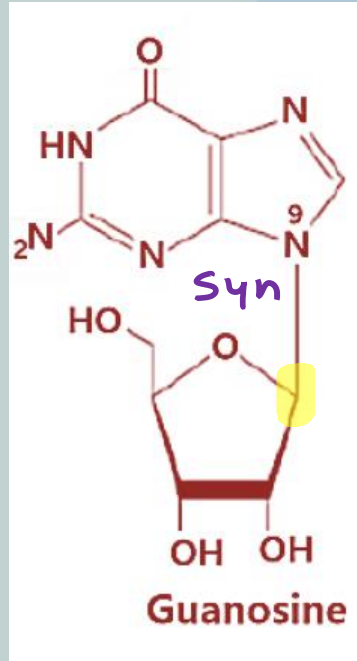
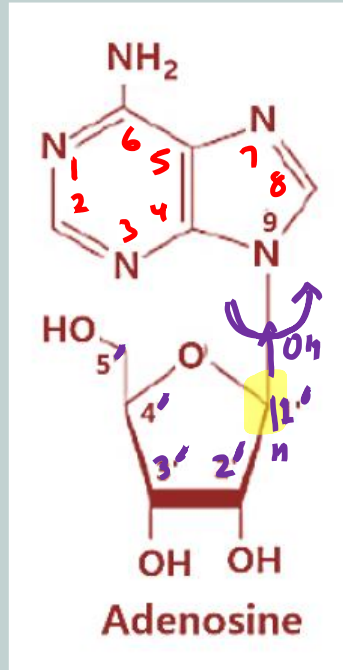


Cytosine (C)

Thymine (T)



Purine Nucleoside : Base + Ribose sugar
Anti/Syn • $\beta 1' \rightarrow N9$ glycosidic bond



Adenine + Ribose =
 Adenosine

Guanine + Ribose =
 Guanosine

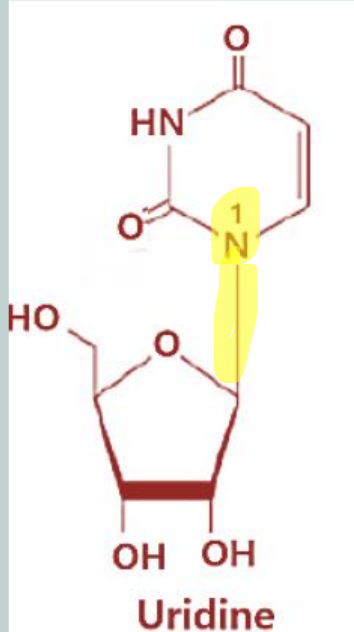
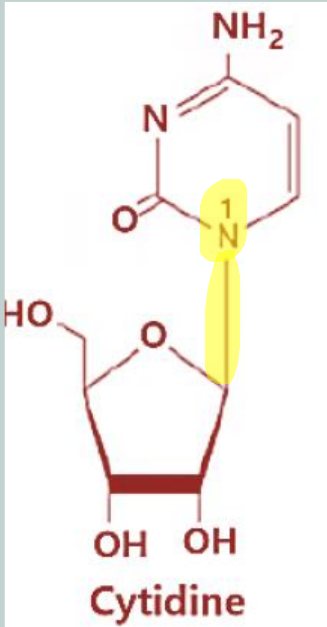
Xanthine + Ribose =
 Xanthosine

Hypoxanthine + Ribose =
 Inosine

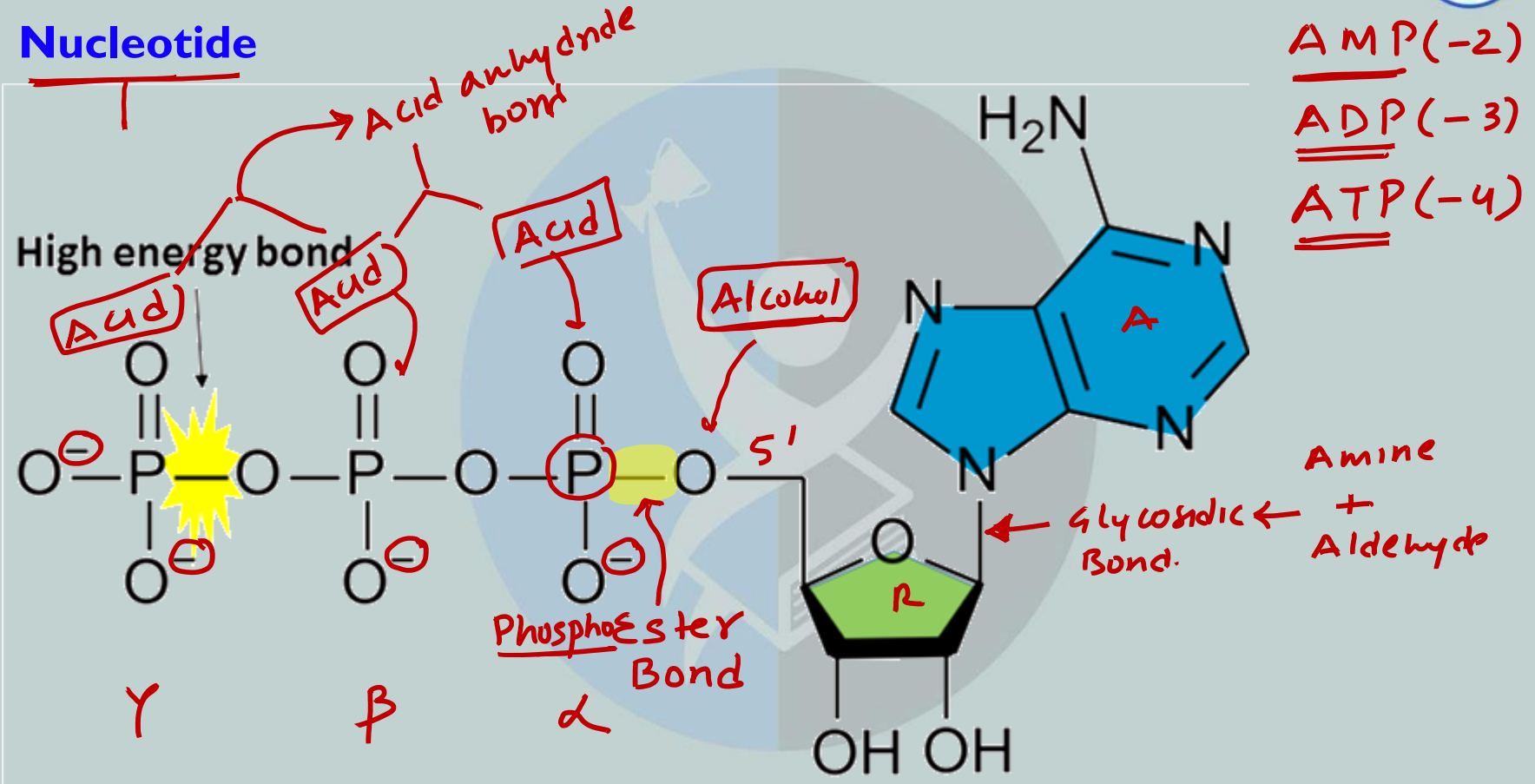


Pyrimidine Nucleoside : β 1' \rightarrow N1 glycosidic bond (Anti/syn)

- Cytosine + Ribose = **Cytidine**
- Uracil + Ribose = **Uridine**
- Thymine + Ribose = **Thymidine**



Nucleotide

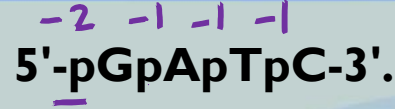


AMP (-2)

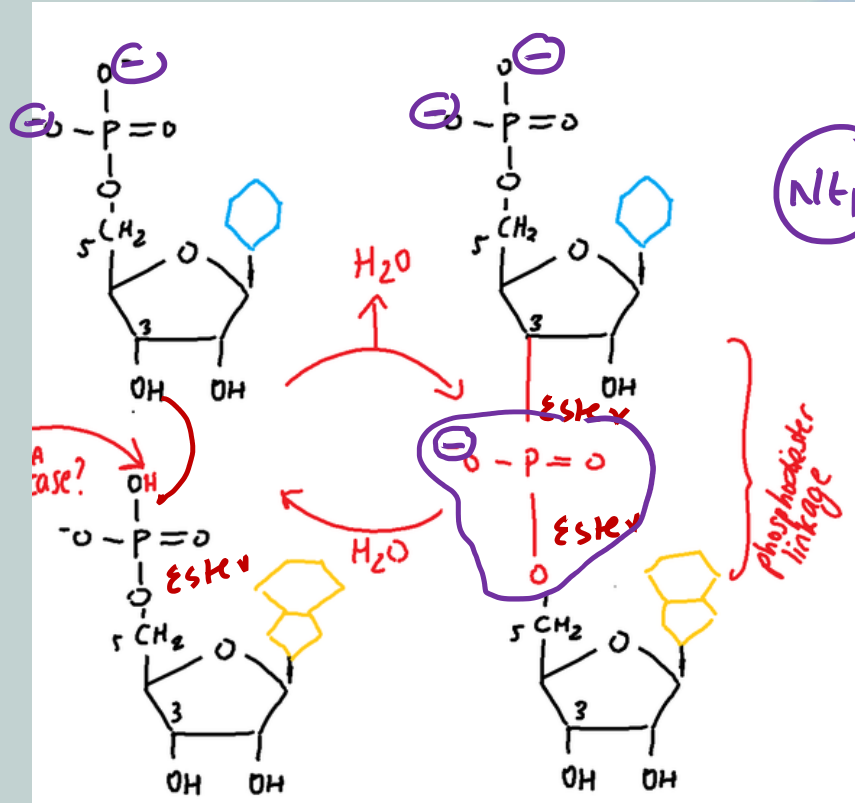
ADP (-3)

ATP (-4)

Polynucleotides:



pn = 7
charge = -5



• Phosphodiester bond

(NT1)



(NT2)

$$\begin{matrix} PPP & Np & Np & Np & N \\ \overline{-4} & \overline{-1} & \overline{-1} & \overline{-1} & \\ = & -7 \end{matrix}$$

→ RNA polymerase
→ DNA polymerase



Average molecular weight

Nucleotide = 330 Dalton (g/mole)

Nucleotide pair (bp) = 660 Dalton (g/mole)

} Part of
nucleic acid

- 100 nt ss RNA, MW = ?
 $100 \times 330 \text{ D} = 33000 \text{ D} = 33 \text{ KD}$
- 100 bp ds DNA, MW ?
 $100 \times 660 \text{ D} = 66000 \text{ D} = 66 \text{ KD}$



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